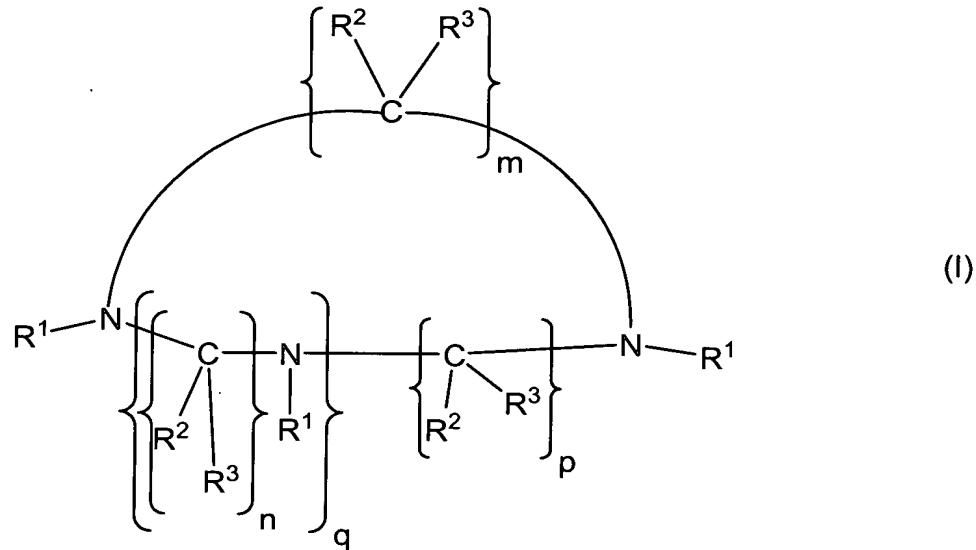


WHAT IS CLAIMED IS:

1 1. A pharmaceutical composition comprising (a) a complex of (i) a
2 cyclic polyaza chelator having complexing affinity for first transition series elements
3 and (ii) a cation of a member selected from the group consisting of calcium and
4 magnesium and (b) a pharmacologically acceptable carrier.

1 2. The pharmaceutical composition of claim 1 in which said cyclic
2 polyaza chelator is a chelator having the formula



3 wherein:

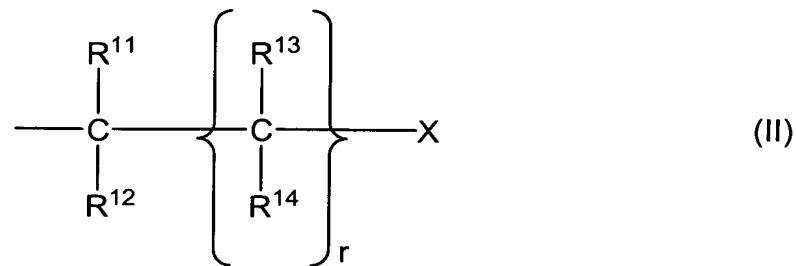
4 5 m, n, and p are each independently 2 or 3;

6 6 q is 1 or 2;

7 7 R² and R³ are each independently selected from the group consisting of H,
8 alkyl, alkenyl, aryl, arylalkyl, alkoxy, alkylthio, alkenoxy, alkenylthio,
9 aryloxy, arylthio, alkyl interrupted by oxa, alkenyl interrupted by oxa,
10 alkyl interrupted by thia, alkenyl interrupted by thia, aryloxyalkyl,
11 alkoxyaryl, aminoalkyl, aminoalkenyl, aminoaryl, aminoarylalkyl,
12 hydroxyalkyl, hydroxyalkenyl, hydroxyaryl, hydroxyarylalkyl, and
13 halogen-substituted versions thereof;

14 14 R¹ is a member selected from the group consisting of R², R³ and radicals of
15 the formula:

16



17

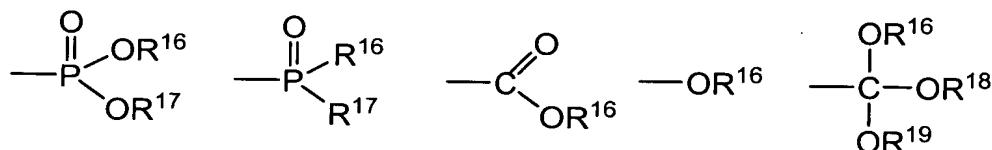
wherein:

18 R^{11} , R^{12} , and R^{13} are each independently selected from the group
 19 consisting of H, alkyl, alkenyl, aryl, arylalkyl, alkoxy, alkylthio,
 20 alkenoxy, alkenylthio, aryloxy, arylthio, alkyl interrupted by oxa,
 21 alkenyl interrupted by oxa, alkyl interrupted by thia, alkenyl
 22 interrupted by thia, aryloxyalkyl, alkoxyaryl, aminoalkyl,
 23 aminoalkenyl, aminoaryl, aminoarylalkyl, hydroxyalkyl,
 24 hydroxyalkenyl, hydroxyaryl, hydroxyarylalkyl, and halogen-
 25 substituted versions thereof;

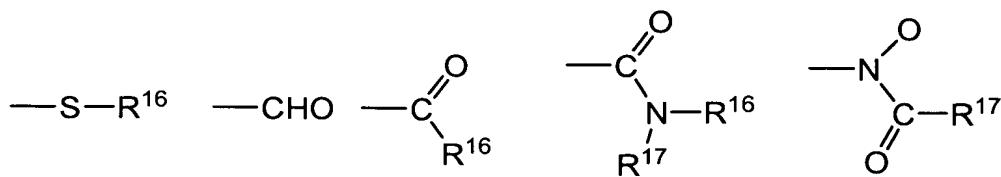
26 R^{14} is a member selected from the group consisting of H, hydroxy,
 27 amino, alkyl, alkyl interrupted by oxa, alkoxy, aryl, aryloxyalkyl,
 28 alkoxyaryl, alkoxyaryl, and halogen-substituted versions thereof;
 29 r is zero or 1; and

30 X is a member selected from the group consisting of alkyl, alkenyl, aryl,
 31 arylalkyl, alkoxy, alkylthio, alkenoxy, alkenylthio, aryloxy,
 32 arylthio, alkyl interrupted by oxa, alkenyl interrupted by oxa, alkyl
 33 interrupted by thia, alkenyl interrupted by thia, aryloxyalkyl,
 34 alkoxyaryl, aminoalkyl, aminoalkenyl, aminoaryl, aminoarylalkyl,
 35 hydroxyalkyl, hydroxyalkenyl, hydroxyaryl, hydroxyarylalkyl,
 36 halogen-substituted versions thereof, and radicals selected from
 37 the group consisting of:

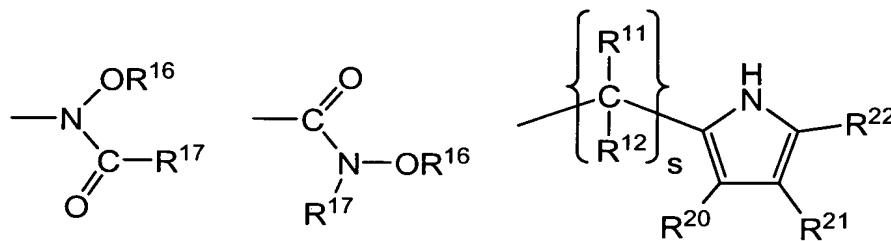
38



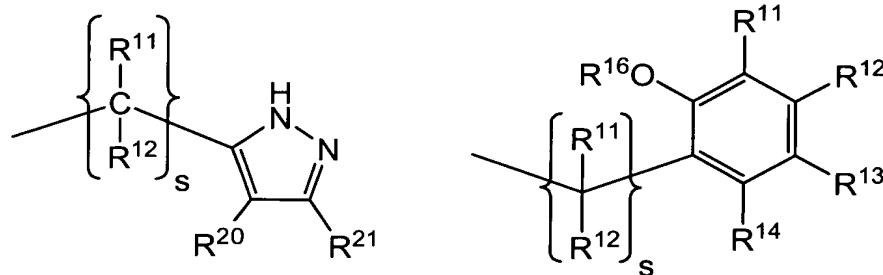
39



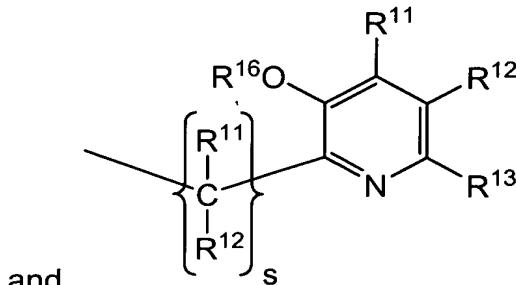
40



41



42



43

wherein,

44

 R^{11} , R^{12} , R^{13} and R^{14} are each independently as defined above;

45

 R^{16} and R^{17} are each independently selected from the group consisting of H, alkyl and aryl, or taken together form a ring structure;

46

 R^{18} and R^{19} are each independently selected from the group consisting of H, alkyl, aryl, alkoxy, alkyl interrupted by oxa, aryloxyalkyl, alkoxyaryl, and halogen-substituted versions thereof;

52

 R^{20} , R^{21} and R^{22} are each independently selected from the group consisting of H, alkyl, alkenyl, aryl, arylalkyl, alkoxy, alkylthio, alkenyloxy, alkenylthio, aryloxy,

53

54

1 3. The pharmaceutical composition of claim 2 wherein m, n, and p
2 are each 2.

4. The pharmaceutical composition of claim 2 wherein q is 1.

1 **5.** The pharmaceutical composition of claim **2** wherein said cation
2 is calcium.

6. The pharmaceutical composition of claim 2 wherein m, n, and p
are each 2, q is 1, and said cation is calcium.

1 7. The pharmaceutical composition of claim 2 wherein all alkyl are
2 C₁-C₄ alkyl.

1 **8.** The pharmaceutical composition of claim **2** wherein all alkyl are
2 C₁-C₄ alkyl, all alkenyl are vinyl, all aryl are phenyl, all aralkyl are phenethyl or
3 benzyl, all cycloalkyl are cyclopentyl or cyclohexyl, and all halogens are chlorine or
4 fluorine.

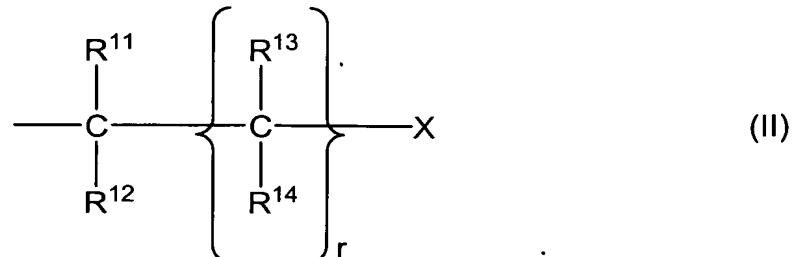
1 9. The pharmaceutical composition of claim 2 wherein R² and R³
2 are each independently selected from the group consisting of H, alkyl, alkenyl, aryl,
3 and aralkyl.

1 **10.** The pharmaceutical composition of claim 2 wherein R² and R³
2 are each independently selected from the group consisting of H and C₁-C₄ alkyl

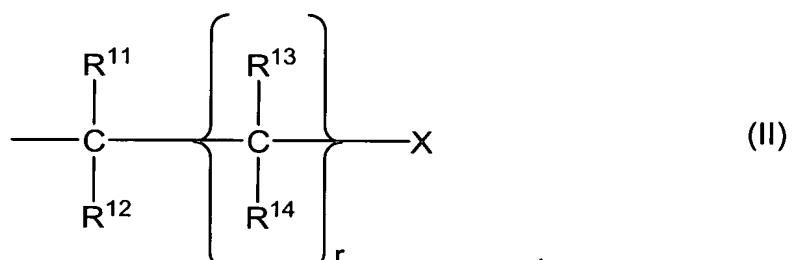
1 **11.** The pharmaceutical composition of claim **2** wherein R² and R³
2 are each H.

1 **12.** The pharmaceutical composition of claim **2** wherein R² and R³
2 are each H and q is 1.

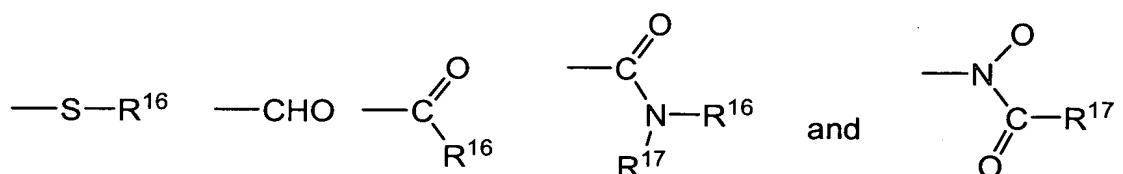
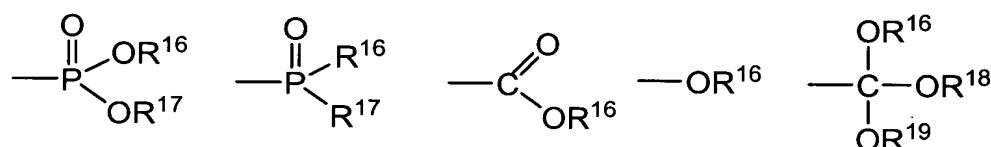
1 **13.** The pharmaceutical composition of claim **2** wherein R¹ is



1 **14.** The pharmaceutical composition of claim **2** wherein q is 1, said
2 cation is calcium, and R¹ is



1 **15.** The pharmaceutical composition of claim **14** wherein X is a
2 member selected from the group consisting of alkyl, alkenyl, aryl, arylalkyl, and
3 radicals selected from the group consisting of:



5
6

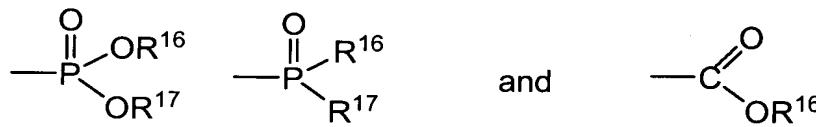
1 **16.** The pharmaceutical composition of claim 15 wherein R¹⁶, R¹⁷,
2 R¹⁸, and R¹⁹ are independently selected from the group consisting of H and C₁-C₄
3 alkyl.

1 **17.** The pharmaceutical composition of claim 14 wherein X is a
2 member selected from the group consisting of alkyl, alkenyl, aryl, arylalkyl, and
3 radicals selected from the group consisting of:



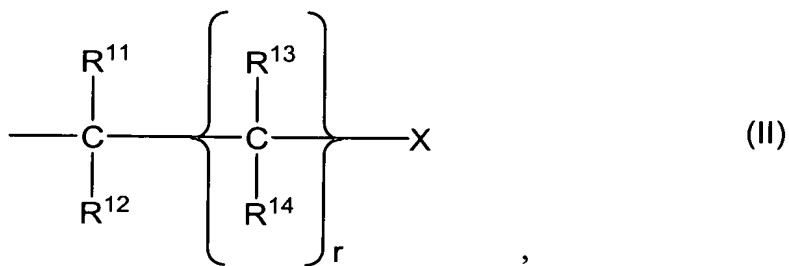
1 **18.** The pharmaceutical composition of claim 17 wherein R¹⁶ and
2 R¹⁷ are independently selected from the group consisting of H and C₁-C₄ alkyl.

1 **19.** The pharmaceutical composition of claim 14 wherein X is a
2 member selected from the group consisting of:



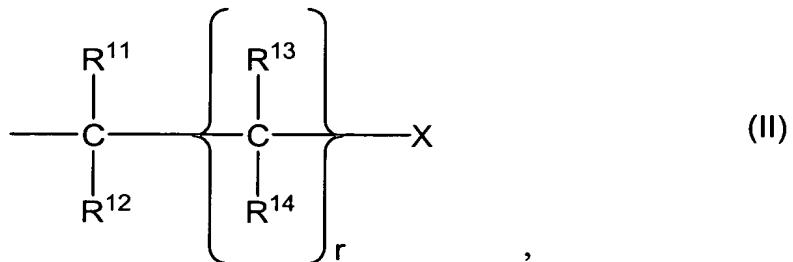
1 **20.** The pharmaceutical composition of claim 19 wherein R¹⁶ and
2 R¹⁷ are independently selected from the group consisting of H and C₁-C₄ alkyl.

1 **21.** The pharmaceutical composition of claim 2 wherein R² and R³
2 are each independently selected from the group consisting of H, alkyl, alkenyl, aryl,
3 and aralkyl, and R¹ is a member selected from the group consisting of H, alkyl,
4 alkenyl, aryl, aralkyl, and



1 in which R¹¹, R¹², and R¹³ are each independently selected from the group consisting
2 of H, alkyl, alkenyl, aryl, and arylalkyl, and R¹⁴ is a member selected from the group
3 consisting of H, hydroxy, amino, and alkyl.

1 **22.** The pharmaceutical composition of claim 2 wherein R¹ is

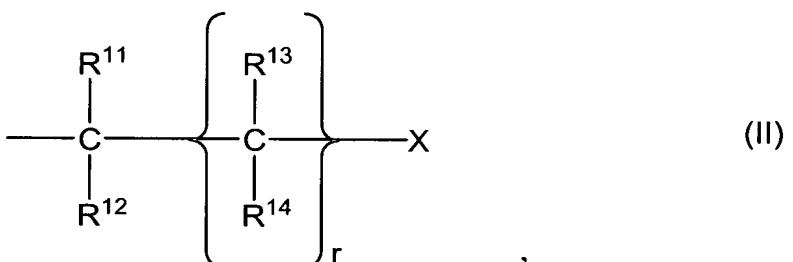


2 ,

1 in which R¹¹, R¹², and R¹³ are each independently selected from the group consisting
2 of H, alkyl, alkenyl, aryl, and arylalkyl, and R¹⁴ is a member selected from the group
3 consisting of H, hydroxy, amino, and alkyl.

1 **23.** The pharmaceutical composition of claim 2 wherein:

2 R¹ is



3 ,

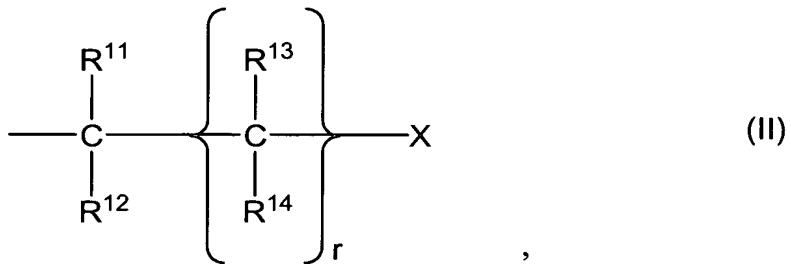
1 in which R¹¹, R¹², and R¹³ are each independently selected from the
2 group consisting of H and C₁-C₄ alkyl, R¹⁴ is a member selected from
3 the group consisting of H and C₁-C₄ alkyl, and X is a member selected
4 from the group consisting of



5 in which R¹⁶ and R¹⁷ are each independently H or C₁-C₄ alkyl;
6 R² and R³ are each independently selected from the group consisting of H
7 and C₁-C₄ alkyl;
8 m, n, and p are each 2;

10 q is 1; and
11 said cation is calcium.

1 **24.** The pharmaceutical composition of claim **2** wherein R¹ is



2 in which R¹¹, R¹², and R¹³ are each independently selected from the group consisting
1 of H and C₁-C₄ alkyl, and R¹⁴ is a member selected from the group consisting of H
2 and C₁-C₄ alkyl.
3

1 **25.** The pharmaceutical composition of claim **2** wherein R¹ is
2 dihydroxyphosphorylmethyl, R² is H, R³ is H, m is 2, n is 2, p is 2, and q is 1.

1 **26.** The pharmaceutical composition of claim **25** in which said cation
2 is calcium.

1 **27.** A method for enhancing the biological activity of a cyclic polyaza
2 chelator having complexing affinity for first transition series elements, said method
3 comprising administering said chelator as a complex with a cation selected from the
4 group consisting of calcium and magnesium.

1 **28.** The method of claim **27** in which said cation is calcium.

2 **29.** A method for providing neuroprotection or cardioprotection in a
3 patient, said method comprising administering to said patient an effective amount of
4 a pharmaceutical composition of claim **1**.

1 **30.** A method for mitigating damage to the central nervous system
2 of a patient suffering from ischemic stroke, seizure or trauma, said method
3 comprising administering to said patient an effective amount of a pharmaceutical
4 composition of claim **1**.

1 **31.** A method for mitigating damage to the heart of a patient
2 suffering a heart attack or arrhythmia, said method comprising administering to said
3 patient an effective amount of a pharmaceutical composition of claim 1.

1 **32.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in a patient, said method comprising administering to said patient an effective
3 amount of a pharmaceutical composition of claim 1.

1 **33.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in a patient that has undergone cardiopulmonary bypass, said method comprising
3 administering to said patient an effective amount of a pharmaceutical composition of
4 claim 1.

1 **34.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in a patient that has undergone vascular surgery, said method comprising
3 administering to said patient an effective amount of a pharmaceutical composition of
4 claim 1.

1 **35.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in transplanted tissue in a patient that has undergone tissue transplant, said method
3 comprising administering to said patient an effective amount of a pharmaceutical
4 composition of claim 1.

1 **36.** A method for providing neuroprotection or cardioprotection in a
2 patient, said method comprising administering to said patient an effective amount of
3 a pharmaceutical composition of claim 2.

1 **37.** A method for enhancing the biological activity of a cyclic polyaza
2 chelator having complexing affinity for first transition series elements, said method
3 comprising administering said chelator as a pharmaceutical composition of claim 2.

1 **38.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in a patient, said method comprising administering to said patient an effective
3 amount of a pharmaceutical composition of claim 2.

1 **39.** A method for mitigating damage to the central nervous system
2 of a patient suffering from ischemic stroke, seizure or trauma, said method
3 comprising administering to said patient an effective amount of a pharmaceutical
4 composition of claim **2**.

1 **40.** A method for mitigating damage to the heart of a patient
2 suffering a heart attack or arrhythmia, said method comprising administering to said
3 patient an effective amount of a pharmaceutical composition of claim **2**.

1 **41.** A method for enhancing the biological activity of a cyclic polyaza
2 chelator having complexing affinity for first transition series elements, said method
3 comprising administering said chelator as a pharmaceutical composition of claim **23**.

1 **42.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in a patient, said method comprising administering to said patient an effective
3 amount of a pharmaceutical composition of claim **23**.

1 **43.** A method for providing neuroprotection or cardioprotection in a
2 patient, said method comprising administering to said patient an effective amount of
3 a pharmaceutical composition of claim **23**.

1 **44.** A method for mitigating damage to the central nervous system
2 of a patient suffering from ischemic stroke, seizure or trauma, said method
3 comprising administering to said patient an effective amount of a pharmaceutical
4 composition of claim **23**.

1 **45.** A method for mitigating damage to the heart of a patient
2 suffering a heart attack or arrhythmia, said method comprising administering to said
3 patient an effective amount of a pharmaceutical composition of claim **23**.

1 **46.** A method for enhancing the biological activity of a cyclic polyaza
2 chelator having complexing affinity for first transition series elements, said method
3 comprising administering said chelator as a pharmaceutical composition of claim **25**.

1 **47.** A method for mitigating ischemia or ischemia-reperfusion injury
2 in a patient, said method comprising administering to said patient an effective
3 amount of a pharmaceutical composition of claim **25**.

1 **48.** A method for providing neuroprotection or cardioprotection in a
2 patient, said method comprising administering to said patient an effective amount of
3 a pharmaceutical composition of claim **25**.

1 **49.** A method for mitigating damage to the central nervous system
2 of a patient suffering from ischemic stroke, seizure or trauma, said method
3 comprising administering to said patient an effective amount of a pharmaceutical
4 composition of claim **25**.

1 **50.** A method for mitigating damage to the heart of a patient
2 suffering a heart attack or arrhythmia, said method comprising administering to said
3 patient an effective amount of a pharmaceutical composition of claim **25**.